

II. Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims

1-75. (Cancelled)

- 76. (Currently amended) An ApoA-I agonist compound comprising:
- (i) an 18 to 22-residue peptide or peptide analogue which forms an amphipathic α -helix in the presence of lipids and which comprises formula (I):

$$Z_{1}-X_{1}-X_{2}-X_{3}-X_{4}-X_{5}-X_{6}-X_{7}-X_{8}-X_{9}-X_{10}-X_{11}-X_{12}-X_{13}-X_{14}-X_{15}-X_{16}-X_{17}-X_{18}-Z_{2}$$

or a pharmaceutically acceptable salt thereof, wherein

X₂ is an aliphatic residue;

 X_3 is Leu (L);

X₄ is an acidic residue;

 X_5 is Leu (L) or Phe (F);

 X_6 is Leu (L) or Phe (F);

 X_7 is a basic residue;

X₈ is an acidic residue;

X₉ is Leu (L) or Trp (W);

 X_{10} is Leu (L) or Trp (W);

 X_{11} is an acidic residue or Asn (N);

 X_{12} is an acidic residue;

 X_{13} is Leu (L), Trp (W) or Phe (F);

 X_{14} is a basic residue or Leu (L);

 X_{15} is Gln (Q) or Asn (N);

 X_{16} is a basic residue;

 X_{17} is Leu (L);

 X_{18} is a basic residue;

wherein at least one <u>L-enantiomeric</u> residue of [the peptide or peptide analogue]

formula (I) is [[a]] replaced with an identical D-enantiomeric residue;

 Z_1 is H_2N_- , or $RC(O)NR_-$;

 Z_2 is -C(O)NRR, -C(O)OR or -C(O)OH;

each R is independently -H, (C_1-C_6) alkyl, (C_1-C_6) alkenyl, (C_1-C_6) alkynyl, (C_5-C_{20}) aryl, (C_6-C_{26}) alkaryl, 5-20 membered heteroaryl or 6-26 membered alkheteroaryl or a 1 to 4-residue peptide or peptide analogue in which one or more bonds between residues 1 through 4 are independently a substituted amide, an isostere of an amide or an amide mimetic;

- each "-" between residues X_1 through X_{18} independently designates an amide linkage, a substituted amide linkage, an isostere of an amide or an amide mimetic;
- (ii) a 14 to 21-residue deleted peptide or peptide analogue according to formula (I) in which at least one and up to eight of residues X_1 , X_2 , X_3 , X_4 , X_5 , X_6 , X_7 , X_8 , X_9 , X_{10} , X_{11} , X_{12} , X_{13} , X_{14} , X_{15} , X_{16} , X_{17} and X_{18} are optionally deleted and wherein at least one <u>remaining L-enantiomeric</u> residue of [the deleted peptide or peptide analogue] <u>formula I</u> is <u>replaced with an identical</u> [[a]] D-enantiomeric residue; or
- (iii) an 18 to 22-residue altered peptide or peptide analogue according to formula (I) in which at least one of residues X_1 , X_2 , X_3 , X_4 , X_5 , X_6 , X_7 , X_8 , X_9 , X_{10} , X_{11} , X_{12} , X_{13} , X_{14} , X_{15} , X_{16} , X_{17} and X_{18} is conservatively substituted and wherein at least one L-enantiomeric residue of the resulting altered peptide or peptide analogue is replaced with an identical [[a]] D-enantiomeric residue; or

an N-terminally blocked form, a C-terminally blocked form or an N- and C-terminally blocked form of formula (I).

- 77. (Canceled)
- 78. (Previously presented) The ApoA-I agonist compound of Claim 76 which is the altered peptide or peptide analogue according to formula (I).
- 79. (Previously presented) The ApoA-I agonist compound of Claim 76 which is the deleted peptide or peptide analogue according to formula (I).
- 80. (Previously presented) The ApoA-I agonist compound of Claim 79 in which one or two helical turns of the peptide or peptide analogue is optionally deleted.

- 81. (Previously presented) The ApoA-I agonist compound of Claim 76 which is an 18-residue peptide or peptide analogue according to formula (I).
- 82. (Previously presented) The ApoA-I agonist compound of Claim 81 in which

the "-" between residues designates -C(O)NH-;

 Z_1 is H_2N_- ; and

 Z_2 is -C(O)OH or a salt thereof.

83. (Previously presented) The ApoA-I agonist compound of Claim 82 in which;

 X_1 is Ala (A), Gly (G), Asn (N) or Pro (P);

X₂ is Ala (A), Val (V) or Leu (L);

 X_3 is Leu (L);

 X_4 is Asp (D) or Glu (E);

 X_5 is Leu (L) or Phe (F);

 X_6 is Leu (L) or Phe (F);

 X_7 is Arg (R), Lys (K) or Orn;

 X_8 is Asp (D) or Glu (E);

 X_9 is Leu (L) or Trp (W);

 X_{10} is Leu (L) or Trp (W);

 X_{11} is Glu (E) or Asn (N);

 X_{12} is Glu (E);

 X_{13} is Leu (L), Trp (W) or Phe (F);

 X_{14} is Arg (R), Lys (K) or Orn;

 X_{15} is Gln (Q) or Asn (N);

 X_{16} is Arg (R), Lys (K) or Orn;

 X_{17} is Leu (L); and

 X_{18} is Arg (R), Lys (K) or Orn.

84. (Currently amended) A multimeric ApoA-I agonist compound which comprises formula (II):

(II) $HH\{LL_m-HH\}_nLL_m-HH$

or a pharmaceutically acceptable salt thereof, wherein:

each m is independently an integer from 0 to 1;

n is an integer from 0 to 10;
each "HH" is independently a peptide or peptide analogue according to
Claim [[1]] 76, the deleted peptide or peptide analogue according to Claim
[[1]] 76 or the altered peptide or peptide analogue according to Claim [[1]] 76;
each "LL" is independently a bifunctional linker; and
each "-" independently designates a covalent linkage; or
an N-terminally blocked form, a C-terminally blocked form or an N- and
C-terminally blocked form of formula (II).

85. (Currently amended) A multimeric ApoA-I agonist compound which comprises formula (III):

(III)
$$X-N_{ya}-X_{(ya-1)}-(N_{yb}-X_{(yb-1)})_p$$

or a pharmaceutically acceptable salt thereof, wherein:

each X is independently [[HHLL_m_HH_nLL_m_HH]] <u>HHfLL_m-HH_nLL_m-HH</u>; each HH is independently a peptide or peptide analogue according to Claim [[1]] <u>76</u>, the deleted peptide or peptide analogue according to Claim [[1]] <u>76</u>; each LL is independently a bifunctional linker; each m is independently an integer from 0 to 1; each n is independently an integer from 0 to 8; N_{ya} and N_{yb} are each independently a multifunctional linking moiety where y_a and y_b represent the number of functional groups on N_{ya} and N_{yb}, respectively; each y_a or y_b is independently an integer from 3 to 8; p is an integer from 0 to 7; and each "—" independently designates a covalent bond; or an N-terminally blocked form, a C-terminally blocked form or an N- and C-terminally blocked form of formula (III).

86. (Currently amended) A multimeric ApoA-I agonist compound which comprises formula (IV) or (V):

or a pharmaceutically acceptable salt thereof, wherein:

each X is independently [[HHLL_m_HH_nLL_m_HH]] <u>HHFLL_m-HH]_nLL_m-HH</u>; each HH is independently a peptide or peptide analogue according to Claim [[1]] <u>76</u>, the deleted peptide or peptide analogue according Claim [[1]] <u>76</u> or the altered peptide or peptide analogue according to Claim [[1]] <u>76</u>;

each LL is independently a bifunctional linker;

each n is independently an integer from 0 to 1;

each m is independently an integer from 0 to 8;

R₁ is -OR or -NRR; and

each R is independently -H, (C1-C6) alkyl, (C1-C6) alkenyl, (C1-C6) alkynyl,

 $(C_5\text{-}C_{20})$ aryl, $(C_6\text{-}C_{26})$ alkaryl, 5-20 membered heteroaryl or 6-26 membered

alkheteroaryl; or

an N-terminally blocked form or a C-terminally blocked form of formula (IV) or (V).

- 87. (Previously presented) The multimeric ApoA-I agonist compound of Claim 84, 85 or 86 in which the bifunctional linker is cleavable.
- 88. (Previously presented) The multimeric ApoA-I agonist compound of Claim 84, 85 or 86 in which n is 0.

- 89. (Previously presented) The multimeric ApoA-I agonist compound of Claim 86 in which m is 0.
- 90. (Previously presented) The multimeric ApoA-I agonist compound of Claim 84, 85 or 86 in which each HH is independently an altered peptide or peptide analogue.
- 91. (Previously presented) The multimeric ApoA-I agonist compound of Claim 84, 85, or 86 in which each HH is independently a deleted peptide or peptide analogue.
- 92. (Previously presented) An ApoA-I agonist compound-lipid complex comprising a lipid and an ApoA-I agonist compound according to Claim 76, 84, 85 or 86.
- 93. (Previously presented) The ApoA-I agonist compound-lipid complex of Claim 92 in which the lipid is sphingomyelin.
- 94. (Previously presented) A pharmaceutical composition comprising a pharmaceutically acceptable carrier, excipient or diluent and an ApoA-I agonist compound according to Claim 76, 84, 85 or 86.
- 95. (Previously presented) A pharmaceutical composition comprising an ApoA-I agonist compound-lipid complex wherein the ApoA-I agonist compound-lipid complex is comprised of an ApoA-I agonist compound according to Claim 76, 84, 85 or 86, a lipid and a pharmaceutically acceptable carrier, excipient or diluent.
- 96. (Previously presented) The pharmaceutical composition of Claim 95 in which the lipid is sphingomyelin.
- 97. (Previously presented) The pharmaceutical composition of Claim 96 which is a lyophilized powder.
- 98. (Previously presented) A method of treating a subject suffering from a disorder associated with dyslipidemia, said method comprising the step of administering to the subject an effective amount of an ApoA-I agonist compound according to Claim 76, 84, 85 or 86.

- 99. (Previously presented) A method of treating a subject suffering from septic shock, said method comprising the step of administering to the subject an effective amount of an ApoA-I agonist compound according to Claim 76, 84, 85 or 86.
- 100. (Previously presented) The method of Claim 98 in which said subject is a human.
- 101. (Previously presented) The method of Claim 99 in which said subject is a human.
- 102. (Previously presented) The method of Claim 98 in which about 0.5 mg/kg to about 100 mg/kg ApoA-I agonist compound is administered to said subject.
- 103. (Previously presented) The method of Claim 99 in which about 0.5 mg/kg to about 100 mg/kg ApoA-I agonist compound is administered to said subject